



APPENDIX A

Claim Amended Shown after Response to Office Action Dated November 16, 2001

- 1. (Amended three times) A method of reducing the growth rate of a tumor, comprising contacting a cell within said tumor with (a) a <u>DNA segment</u> [gene] encoding a functional p53 protein and (b) a DNA damaging agent in a combined amount effective to inhibit the growth of said tumor, wherein functional p53 protein is expressed in the cell.
- 2. (Amended three times) The method of claim 1, wherein the DNA damaging agent is [said cell is contacted with said gene in combination with] X-ray radiation, UV-irradiation, γ-irradiation, microwaves, adriamycin, 5-fluorouracil, etoposide, camptothecin, actinomycin-D, mitomycin C, or cisplatin.
- 3. (Amended twice) The method of claim 2, wherein said cell is contacted with the DNA segment [said gene] in combination with cisplatin.
- 4. (Amended three times) The method of claim 1, wherein the DNA segment is in [said cell is contacted with] a recombinant vector that expresses the [a] functional p53 protein in said cell [in combination with a DNA damaging agent].
- 5. (Amended four times) The method of claim 4, wherein said p53-expressing recombinant[,] vector is a naked DNA plasmid, [or] a plasmid within a liposome, a retroviral vector, an AAV vector, or a recombinant adenoviral vector.
- 8. (Amended three times) The method of claim 4, wherein said recombinant vector comprises a p53 expression region, <u>a</u> [the] cytomegalovirus IE promoter and <u>an</u> [the] SV40 early polyadenylation signal.



- 9. (Amended) The method of claim 6, wherein at least one gene essential for adenovirus replication is deleted from said adenovirus vector [construct] and a [the] p53 expression region is introduced in its place.
- 10. (Amended) The method of claim 9, wherein [the] E1A and E1B regions of the adenovirus vector are deleted and the p53 expression region is introduced in their place.
- 11. (Cancelled)
- 12. (Amended twice) The method of claim 1, wherein said cell is first contacted with the DNA segment [said gene] and is subsequently contacted with said DNA damaging agent.
- 13. (Amended twice) The method of claim 1, wherein said cell is first contacted with said DNA damaging agent and is subsequently contacted with the DNA segment [said gene].
- 14. (Amended twice) The method of claim 1, wherein said cell is simultaneously contacted with the DNA segment [said gene] and said DNA damaging agent.
- 15. (Amended twice) The method of claim 1, wherein said cell is contacted with a first composition comprising the DNA segment [said gene] and a second composition comprising said DNA damaging agent.
- 17. (Amended twice) The method of claim 1, wherein said cell is contacted with a single composition comprising the DNA segment [said gene] in combination with said DNA damaging agent.
- 22. (Amended three times) The method of claim 1, wherein said [tumor] cell is a malignant cell.



- 26. (Amended four times) The method of claim 1, wherein said [tumor] cell is located within an animal at a tumor site.
- 32. (Amended twice) A composition comprising <u>a) an exogenous DNA segment</u> [a gene] encoding a functional p53 polypeptide <u>and b)</u> [in combination with] a DNA damaging agent.
- 33. (Amended three times) The composition of claim 32, wherein the DNA damaging agent is [comprising said gene in combination with] adriamycin, 5-fluorouracil, etoposide, camptothecin, actinomycin-D, mitomycin C, or cisplatin.
- 34. (Amended twice) The composition of claim 33, wherein the DNA damaging agent is [comprising said gene in combination with] cisplatin.
- 35. (Amended twice) The composition of claim 32, wherein the exogenous DNA segment is in [comprising] a recombinant vector that expresses a functional p53 protein in an animal cell [in combination with a DNA damaging agent].
- 38. (Cancelled)
- 39. (Amended) The composition of claim 37, wherein the recombinant vector is [32, comprising] a recombinant adenoviral vector and the DNA damaging agent is [present within a recombinant adenovirus particle in combination with] cisplatin.
- 45. (Amended) The kit of claim 42, wherein the recombinant vector is an [comprising a pharmaceutical formulation of a recombinant] adenovirus [including a recombinant] vector [that expresses a p53 protein in an animal cell] and the DNA damaging agent is [a pharmaceutical formulation of] cisplatin.





- 46. (Amended twice) The method of claim 1, wherein the [tumor] cell is contacted with a DNA damaging agent by irradiating the [tumor] cell with X-ray radiation, UV-irradiation, γ-irradiation or microwaves.
- 47. (Amended) The method of claim 46, wherein the [tumor] cell is contacted with a DNA damaging agent by irradiating the [tumor] cell with X-ray radiation.
- 48. (Amended) The method of claim 46, wherein the [tumor] cell is contacted with a DNA damaging agent by irradiating the [tumor] cell with UV-irradiation.
- 49. (Amended) The method of claim 46, wherein the [tumor] cell is contacted with a DNA damaging agent by irradiating the [tumor] cell with γ-irradiation.
- 50. (Amended) The method of claim 46, wherein the [tumor] cell is contacted with a DNA damaging agent by irradiating the [tumor] cell with microwaves.
- 51. (Amended twice) The method claim 1, wherein the [tumor] cell is contacted with a pharmaceutical composition comprising the [a] DNA damaging agent [compound].
- 77. (Amended twice) The method of claim 4, wherein said <u>DNA segment</u> [gene] is administered prior to said DNA damaging agent.
- 78. (Amended twice) The method of claim 4, wherein said <u>DNA segment</u> [gene] is administered after said DNA damaging agent.
- 79. (Amended twice) The method of claim 4, wherein said <u>DNA segment</u> [gene] is administered at the same time as said DNA damaging agent.



- 83. (Amended three times) The method of claim 26, wherein said <u>DNA segment</u> [gene] is delivered to said tumor endoscopically, intravenously, intratracheally, intralesionally, percutaneously or subcutaneously.
- 86. (Amended twice) The method of claim 13, wherein there is 12 to 24 hours [the period] between administration of the DNA damaging agent and administration of the DNA segment [gene is between 12 and 24 hours].
- 87. (Amended twice) The method of claim 13, wherein there is 6 to 12 hours [the period] between administration of the DNA damaging agent and administration of the DNA segment [gene is between 6 and 12 hours].
- 88. (Amended twice) The method of claim 13, wherein there is about 12 hours [the period] between administration of the DNA damaging agent and administration of the DNA segment [gene is about 12 hours].
- 89. (Amended twice) The method of claim 12, wherein there is 12 to 24 hours [the period] between administration of the DNA segment [gene] and administration of the DNA damaging agent [is between 12 and 24 hours].
- 90. (Amended three times) The method of claim 12, wherein there is 6 to 12 hours [the period] between administration of the DNA segment [gene] and administration of the DNA damaging agent [is between 6 and 12 hours].
- 91. (Amended three times) The method of claim 12, wherein there is about 12 hours [the period] between administration of the DNA segment [gene] and administration of the DNA damaging agent [is about 12 hours].
- 101. (Amended) The method of claim 23 [95], wherein said lung cancer cell is a small cell lung carcinoma cell.



- 119. (Amended) The method of claim 47, wherein the <u>cell is irradiated with about [x-ray dosage is between] 2000 to [and] 6000 roentgens.</u>
- 120. (Amended) The method of claim 47, wherein the <u>cell is irradiated with about [x-ray dosage is between] 50 to [and] 200 roentgens.</u>